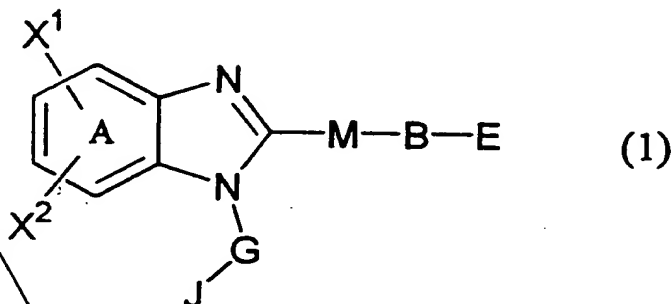


CLAIMS

1. An inhibitor against human chymase activity containing a benzimidazole derivative expressed by the following formula (1) or its salt as an active ingredient,



[in the formula (1), the ring marked with A expresses a pyridine ring or a benzene ring;

X<sup>1</sup> and X<sup>2</sup> are each at the same time or independently a hydrogen atom, a halogen atom, a trihalomethyl group, a hydroxyl group, a nitro group, a cyano group, -CH<sub>2</sub>NH<sub>2</sub>, -CH=NR<sup>1</sup>, -CH=NOR<sup>1</sup> or -CONR<sup>1</sup>R<sup>2</sup> (here, R<sup>1</sup> and R<sup>2</sup> are each a hydrogen atom or a C<sub>1-4</sub> alkyl group), -COOR<sup>3</sup> (here, R<sup>3</sup> is a hydrogen atom or a C<sub>1-4</sub> alkyl group), a substituted or unsubstituted C<sub>1-6</sub> normal, cyclic or branched alkyl group, a substituted or unsubstituted C<sub>3-7</sub> cycloalkyl group, a substituted or unsubstituted C<sub>1-6</sub> normal or branched alkoxy group, a substituted or unsubstituted C<sub>1-6</sub> normal or branched alkylthio group, a substituted or unsubstituted C<sub>1-6</sub> normal or branched alkylsulfonyl group or a substituted or unsubstituted C<sub>1-6</sub> normal or branched alkylsulfinyl group (the substituent permissible to the groups is a halogen atom, a hydroxyl group, a nitro group, a cyano group, an acyl group, a trihalomethyl group, a trihalomethoxy group, a phenyl group, an oxo group or a phenoxy group optionally substituted with one or more halogen atoms, and the substituent may substitute singly or plurally independently at arbitrary position(s));

B is a substituted or unsubstituted C<sub>1-6</sub> normal, cyclic or branched alkylene group or a substituted or unsubstituted C<sub>2-6</sub> normal or branched alkenylene group (the substituent permissible to the groups is a halogen atom, a hydroxyl group, a nitro group, a cyano group, a C<sub>1-6</sub> normal or branched alkoxy

group (including the case where adjacent two groups form an acetal bonding), a C<sub>1-6</sub> normal or branched alkylthio group, a C<sub>1-6</sub> normal or branched alkylsulfonyl group, a C<sub>1-6</sub> normal or branched acyl group, a C<sub>1-6</sub> normal or branched acylamino group, a trihalomethyl group, a trihalomethoxy group, a phenyl group, an oxo group or a phenoxy group optionally substituted with one or more halogen atoms, and the substituent may substitute singly or plurally independently at arbitrary position(s) of the alkylene group or an alkenylene group; between atoms, the alkylene group or alkenylene group optionally contains one or more of -O-, -S-, -SO<sub>2</sub>- or -NR<sup>4</sup>-, but this atom or atomic group does not bond directly to the M, and here R<sup>4</sup> is a hydrogen atom or a C<sub>1-6</sub> normal or branched alkyl group};

E expresses -COOR<sup>4</sup>, -SO<sub>3</sub>R<sup>4</sup>, -CONHR<sup>5</sup>, -SO<sub>2</sub>NHR<sup>4</sup>, -PO(OR<sup>6</sup>)<sub>2</sub>, a tetrazol-5-yl group, a 5-oxo-1,2,4-oxadiazol-3-yl group or a 5-oxo-1,2,4-thiadiazol-3-yl group (here, R<sup>4</sup> is similarly defined as above; R<sup>5</sup> is a hydrogen atom, a cyano group, or a C<sub>1-6</sub> normal or branched alkyl group; R<sup>6</sup> is a hydrogen atom, a C<sub>1-6</sub> normal or branched alkyl group, or trifluoromethylsulfonyl group, or its pharmaceutically permissible salt);

G is a substituted or unsubstituted C<sub>1-6</sub> normal or branched alkylene group (between atoms, the alkylene group optionally contains one or more of -O-, -S-, -SO<sub>2</sub>- or -NR<sup>4</sup>-, but this atom or atomic group does not bond directly to the nitrogen atom of the imidazole ring (R<sup>4</sup> is similarly defined as above), and the substituent is a halogen atom, a hydroxyl group, a nitro group, a cyano group, a C<sub>1-6</sub> normal or branched alkoxyl group (including the case where adjacent two groups form an acetal bonding), a trihalomethyl group, a trihalomethoxy group, a phenyl group or an oxo group};

J is a substituted or unsubstituted C<sub>1-6</sub> normal, cyclic or branched alkyl group, a substituted or unsubstituted C<sub>4-10</sub> aryl group (the substituent permissible to the groups is a halogen atom, a hydroxyl group, a nitro group, a cyano group, -COOR<sup>7</sup> (here, R<sup>7</sup> is a hydrogen atom or a C<sub>1-4</sub> alkyl group), a C<sub>1-6</sub> normal, cyclic or branched alkyl group, a C<sub>1-6</sub> normal or branched alkoxyl group (including the case where adjacent two groups form an acetal bonding), a C<sub>1-6</sub> normal or branched alkylthio group, a C<sub>1-6</sub> normal or branched alkylsulfonyl group, a C<sub>1-6</sub> normal or branched alkylsulfinyl group, a C<sub>1-6</sub> acyl group, a C<sub>1-6</sub> normal or branched acylamino group, a trihalomethyl group, a trihalomethoxy

group, a phenyl group, an oxo group, or a phenoxy group optionally substituted with one or more halogen atoms; the substituent may substitute singly or plurally independently at arbitrary position(s) of the alkyl group or aryl group; and the substituent is further optionally substituted with a halogen atom, a hydroxyl group, a nitro group, a cyano group, an acyl group, a trihalomethyl group, a phenyl group, an oxo group or a phenoxy group optionally substituted with a halogen atom); and

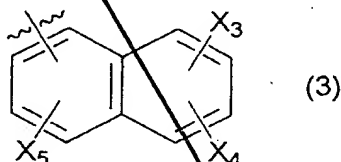
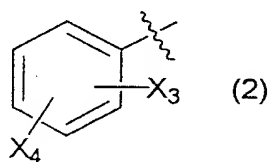
M is a sulfur atom, a sulfinyl group, a sulfonyl group, a single bond or  $-CR^8R^9-$  (here,  $R^8$  and  $R^9$  are each at the same time or independently a hydrogen atom or a  $C_{1-4}$  alkyl group)].

2. An inhibitor against human chymase activity set forth in Claim 1 wherein the ring marked with A in the above formula (1) is a benzene ring.

3. An inhibitor against human chymase activity set forth in Claim 1 wherein the ring marked with A in the above formula (1) is a pyridine ring.

4. An inhibitor against human chymase activity set forth in one out of Claims 1 to 3 wherein  $X^1$  and  $X^2$  in the above formula (1) are each at the same time or independently a hydrogen atom, a halogen atom, a trihalomethyl group, a cyano group, a substituted or unsubstituted  $C_{1-3}$  normal or branched alkyl group, a substituted or unsubstituted  $C_{1-3}$  normal or branched alkoxy group, or a substituted or unsubstituted  $C_{1-3}$  normal or branched alkylthio group.

5. An inhibitor against human chymase activity set forth in one out of Claims 1 to 4 wherein J in the above formula (1) is a group described in the following formula (2) or (3),



[here,  $X^3$ ,  $X^4$  and  $X^5$  are each at the same time or independently a hydrogen atom, a halogen atom, a hydroxyl group, a nitro group, a cyano group, a trihalomethyl group, a trihalomethoxy group,  $-COOR^7$  (here,  $R^7$  is a hydrogen atom or a  $C_{1-4}$  alkyl group), a substituted or unsubstituted  $C_{1-3}$  normal or branched alkyl group, a substituted or unsubstituted  $C_{1-3}$  normal or branched

alkoxyl group, a substituted or unsubstituted C<sub>1-3</sub> normal or branched alkylthio group, a substituted or unsubstituted C<sub>1-3</sub> normal or branched alkylsulfonyl group, or a substituted or unsubstituted C<sub>1-3</sub> normal or branched alkylsulfinyl group; there is no limitation regarding the substitution positions of X<sup>3</sup>, X<sup>4</sup> and X<sup>5</sup> on the benzene ring or the naphthalene ring].

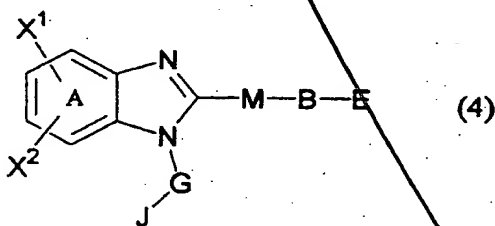
6. An inhibitor against human chymase activity set forth in one out of Claims 1 to 5 wherein M in the above-mentioned formula (1) is a sulfur atom.

7. An inhibitor against human chymase activity set forth in one out of Claims 1 to 6 wherein B in the above-mentioned formula (1) is a substituted or unsubstituted C<sub>1-6</sub> normal, cyclic or branched alkylene group.

8. An inhibitor against human chymase activity set forth in one out of Claims 1 to 7 wherein G in the above-mentioned formula (1) is -CH<sub>2</sub>-, -CH<sub>2</sub>CH<sub>2</sub>-, -CH<sub>2</sub>CO-, -CH<sub>2</sub>CH<sub>2</sub>O-, -CH<sub>2</sub>CONH-, -CO-, -SO<sub>2</sub>-, -CH<sub>2</sub>SO<sub>2</sub>-, -CH<sub>2</sub>S- or -CH<sub>2</sub>CH<sub>2</sub>S- (J bonds to the right side of said group).

9. An inhibitor against human chymase activity set forth in one out of Claims 1 to 8 wherein E in the above-mentioned formula (1) is -COOH.

10. A benzimidazole derivative expressed by the following formula (4) or its pharmaceutically permissible salt,



[in the formula (4), the definitions of the ring marked with A, and X<sup>1</sup>, X<sup>2</sup>, B, E, G, J and M are same as those in the above formula (1); however, excepting the case where at least one of X<sup>1</sup> and X<sup>2</sup> is a cyano group, -CH<sub>2</sub>NH<sub>2</sub>, -CH=NR<sup>1</sup>, -CH=NOR<sup>1</sup> or -CONR<sup>1</sup>R<sup>2</sup> (here, R<sup>1</sup> and R<sup>2</sup> are each a hydrogen atom or a C<sub>1-4</sub> alkyl group), J expresses only a substituted naphthalene ring].

11. A benzimidazole derivative or its pharmaceutically permissible salt set forth in Claim 10 wherein X<sup>1</sup> and X<sup>2</sup> in the above formula (4) are each a hydrogen atom, a cyano group, -CH<sub>2</sub>NH<sub>2</sub>, -CH=NR<sup>1</sup>, -CH=NOR<sup>1</sup> or -CONR<sup>1</sup>R<sup>2</sup> (here, R<sup>1</sup> and R<sup>2</sup> are each a hydrogen atom or a C<sub>1-4</sub> alkyl group; X<sup>1</sup> and X<sup>2</sup> are not hydrogen at the same time).

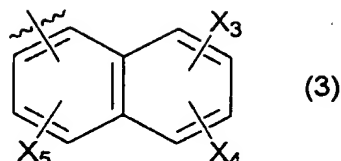
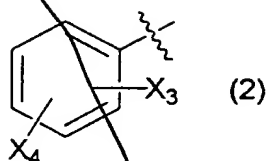
12. A benzimidazole derivative or its pharmaceutically permissible salt set forth in Claim 10 wherein  $X^1$  and  $X^2$  in the above formula (4) are each at the same time or independently a hydrogen atom, a halogen atom, a trihalomethyl group, a hydroxyl group, a nitro group,  $-CH=NR^1$  (here,  $R^1$  is a hydrogen atom or a  $C_{1-4}$  alkyl group),  $-COOR^3$  (here,  $R^3$  is a hydrogen atom or a  $C_{1-4}$  alkyl group), a substituted or unsubstituted  $C_{1-6}$  normal, cyclic or branched alkyl group, a substituted or unsubstituted  $C_{3-7}$  cycloalkyl, a substituted or unsubstituted  $C_{1-6}$  normal or branched alkoxy group, a substituted or unsubstituted  $C_{1-6}$  normal or branched alkylthio group, a substituted or unsubstituted  $C_{1-6}$  normal or branched alkylsulfonyl group or a substituted or unsubstituted  $C_{1-6}$  normal or branched alkylsulfinyl group {the substituent permissible to the groups is a halogen atom, a hydroxyl group, a nitro group, a cyano group, an acyl group, a trihalomethyl group, a trihalomethoxy group, a phenyl group, an oxo group or a phenoxy group optionally substituted with one or more halogen atoms, and the substituent may substitute singly or plurally independently at arbitrary position(s)}.

13. A benzimidazole derivative or its pharmaceutically permissible salt set forth in Claim 10 wherein  $X^1$  and  $X^2$  in the above formula (4) are each a hydrogen atom or a cyano group (here,  $X^1$  and  $X^2$  can not be hydrogen atoms at the same time).

14. A benzimidazole derivative or its pharmaceutically permissible salt set forth in one out of Claims 10 to 13 wherein M in the above formula (4) is a sulfur atom.

15. A benzimidazole derivative or its pharmaceutically permissible salt set forth in one out of Claims 10 to 14 wherein B in the above formula (4) is a substituted or unsubstituted  $C_{1-6}$  normal, cyclic or branched alkylene group.

16. A benzimidazole derivative or its pharmaceutically permissible salt set forth in one out of Claims 10 to 15 wherein J in the above formula (4) is a group expressed by the following formula (2) or (3),



[here, X<sup>3</sup>, X<sup>4</sup> and X<sup>5</sup> are each at the same time or independently a hydrogen atom, a halogen atom, a hydroxyl group, a nitro group, a cyano group, a trihalomethyl group, a trihalomethoxy group, -COOR<sup>7</sup> (here, R<sup>7</sup> is a hydrogen atom or a C<sub>1-4</sub> alkyl group), a substituted or unsubstituted C<sub>1-3</sub> normal or branched alkyl group, a substituted or unsubstituted C<sub>1-3</sub> normal or branched alkoxyl group, a substituted or unsubstituted C<sub>1-3</sub> normal or branched alkylthio group, a substituted or unsubstituted C<sub>1-3</sub> normal or branched alkylsulfonyl group, or a substituted or unsubstituted C<sub>1-3</sub> normal or branched alkylsulfinyl group; there is no limitation regarding the substitution positions of X<sup>3</sup>, X<sup>4</sup> and X<sup>5</sup> on the benzene ring or the naphthalene ring].

17. A benzimidazole derivative or its pharmaceutically permissible salt set forth in one out of Claims 10 to 16 wherein G in the above formula (4) is -CH<sub>2</sub>-, -CH<sub>2</sub>CH<sub>2</sub>-, -CH<sub>2</sub>CO-, -CH<sub>2</sub>CH<sub>2</sub>O-, -CH<sub>2</sub>CONH-, -CO-, -SO<sub>2</sub>-, -CH<sub>2</sub>SO<sub>2</sub>-, -CH<sub>2</sub>S- or -CH<sub>2</sub>CH<sub>2</sub>S- (J bonds to the right side of said group).

18. A benzimidazole derivative or its pharmaceutically permissible salt set forth in one out of Claims 10 to 17 wherein E in the above formula (4) is COOH.

19. A pharmaceutical composition consisting of a benzimidazole derivative and/or its pharmaceutically permissible salt set forth in one out of Claims 10 to 18, and a pharmaceutically permissible carrier.

20. A chymase activity inhibitor set forth in one out of Claims 1 to 9 whose targeting disease is an inflammatory disease, an allergy disease, a respiratory disease, a cardiovascular disease or a bone/cartridge metabolic disease.

21. A human chymase activity inhibitor set forth in Claim 20 which is a preventing agent or a treating agent of a disease.